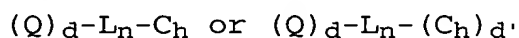


WHAT IS CLAIMED IS:

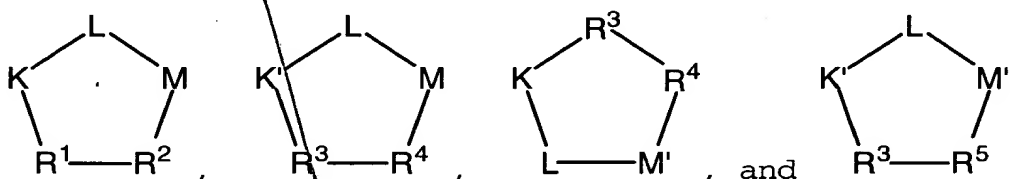
1. A compound, comprising: a targeting moiety and a chelator, wherein the targeting moiety is bound to the chelator, is a peptide or peptidomimetic, and binds to a receptor that is upregulated during angiogenesis and the compound has 0-1 linking groups between the targeting moiety and chelator.

2. A compound according to Claim 1, wherein the targeting moiety is a peptide or a mimetic thereof and the receptor is selected from the group: EGFR, FGFR, PDGFR, Flk-1/KDR, Flt-1, Tek, Tie, neuropilin-1, endoglin, endosialin, Axl, $\alpha_v\beta_3$, $\alpha_v\beta_5$, $\alpha_5\beta_1$, $\alpha_4\beta_1$, $\alpha_1\beta_1$, and $\alpha_2\beta_2$ and the linking group is present between the targeting moiety and chelator.

3. A compound according to Claim 2, the receptor is the integrin $\alpha_v\beta_3$ and the compound is of the formula:



wherein, Q is a peptide independently selected from the group:



K is an L-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine, δ -N-2-imidazolinylnornithine, δ -N-benzylcarbamoynornithine, and β -2-benzimidazolylacetyl-1,2-diaminopropionic acid;

K' is a D-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine,

δ -N-2-imidazolinylnornithine,
 δ -N-benzylcarbamoylnornithine, and
 β -2-benzimidazolylacetyl-1,2-diaminopropionic acid;

5 L is independently selected at each occurrence from the group:
glycine, L-alanine, and D-alanine;

M is L-aspartic acid;

10 M' is D-aspartic acid;

Sub
BI
cont
15 R¹ is an amino acid substituted with 0-1 bonds to L_n,
independently selected at each occurrence from the group:
glycine, L-valine, D-valine, alanine, leucine,
isoleucine, norleucine, 2-aminobutyric acid,
2-aminohexanoic acid, tyrosine, phenylalanine,
thienylalanine, phenylglycine, cyclohexylalanine,
homophenylalanine, 1-naphthylalanine, lysine, serine,
20 ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic
acid, cysteine, penicillamine, and methionine;

R² is an amino acid, substituted with 0-1 bonds to L_n,
independently selected at each occurrence from the group:
glycine, valine, alanine, leucine, isoleucine,
25 norleucine, 2-aminobutyric acid, 2-aminohexanoic acid,
tyrosine, L-phenylalanine, D-phenylalanine,
thienylalanine, phenylglycine, biphenylglycine,
cyclohexylalanine, homophenylalanine,
L-1-naphthylalanine, D-1-naphthylalanine, lysine, serine,
30 ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic
acid, cysteine, penicillamine, methionine, and
2-aminothiazole-4-acetic acid;

R³ is an amino acid, substituted with 0-1 bonds to L_n,
35 independently selected at each occurrence from the group:
glycine, D-valine, D-alanine, D-leucine, D-isoleucine,
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic
acid, D-tyrosine, D-phenylalanine, D-thienylalanine,

000000 448260

Sub
31
cont

5 D-phenylglycine, D-cyclohexylalanine,
D-homophenylalanine, D-1-naphthylalanine, D-lysine,
D-serine, D-ornithine, D-1,2-diaminobutyric acid,
D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine,
and D-methionine;

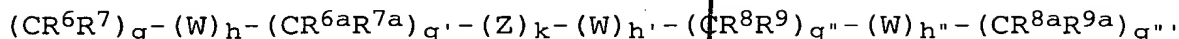
10 R⁴ is an amino acid, substituted with 0-1 bonds to L_n,
independently selected at each occurrence from the group:
glycine, D-valine, D-alanine, D-leucine, D-isoleucine,
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic
acid, D-tyrosine, D-phenylalanine, D-thienylalanine,
D-phenylglycine, D-cyclohexylalanine,
D-homophenylalanine, D-1-naphthylalanine, D-lysine,
D-serine, D-ornithine, D-1,2-diaminobutyric acid,
D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine,
D-methionine, and 2-aminothiazole-4-acetic acid;

20 R⁵ is an amino acid, substituted with 0-1 bonds to L_n,
independently selected at each occurrence from the group:
glycine, L-valine, L-alanine, L-leucine, L-isoleucine,
L-norleucine, L-2-aminobutyric acid, L-2-aminohexanoic
acid, L-tyrosine, L-phenylalanine, L-thienylalanine,
L-phenylglycine, L-cyclohexylalanine,
L-homophenylalanine, L-1-naphthylalanine, L-lysine,
25 L-serine, L-ornithine, L-1,2-diaminobutyric acid,
L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine,
L-methionine, and 2-aminothiazole-4-acetic acid;

30 provided that one of R¹, R², R³, R⁴, and R⁵ in each Q is
substituted with a bond to L_n, further provided that when
R² is 2-aminothiazole-4-acetic acid, K is
N-methylarginine, further provided that when R⁴ is
2-aminothiazole-4-acetic acid, K and K' are
N-methylarginine, and still further provided that when R⁵
35 is 2-aminothiazole-4-acetic acid, K' is N-methylarginine;

d is selected from 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

L_n is a linking group having the formula:



5 provided that $g+h+g'+k+h'+g''+h''+g'''$ is other than 0;

W is independently selected at each occurrence from the group:

10 O, S, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O),
NHC(=S)NH, NHC(=O)NH, SO₂, (OCH₂CH₂)_s, (CH₂CH₂O)_{s'},
(OCH₂CH₂CH₂)_{s''}, (CH₂CH₂CH₂O)_t, and (aa)_{t'};

aa is independently at each occurrence an amino acid;

15 Z is selected from the group: aryl substituted with 0-3 R¹⁰,
C₃₋₁₀ cycloalkyl substituted with 0-3 R¹⁰, and a 5-10
membered heterocyclic ring system containing 1-4
heteroatoms independently selected from N, S, and O and
substituted with 0-3 R¹⁰;

20 R⁶, R^{6a}, R⁷, R^{7a}, R⁸, R^{8a}, R⁹ and R^{9a} are independently selected
at each occurrence from the group: H, =O, COOH, SO₃H,
PO₃H, C₁₋₅ alkyl substituted with 0-3 R¹⁰, aryl
substituted with 0-3 R¹⁰, benzyl substituted with 0-3
R¹⁰, and C₁₋₅ alkoxy substituted with 0-3 R¹⁰,
25 NHC(=O)R¹¹, C(=O)NHR¹¹, NHC(=O)NHR¹¹, NHR¹¹, R¹¹, and a
bond to C_H;

30 R¹⁰ is independently selected at each occurrence from the
group: a bond to C_H, COOR¹¹, OH, NHR¹¹, SO₃H, PO₃H, aryl
substituted with 0-3 R¹¹, C₁₋₅ alkyl substituted with 0-1
R¹², C₁₋₅ alkoxy substituted with 0-1 R¹², and a 5-10
membered heterocyclic ring system containing 1-4
heteroatoms independently selected from N, S, and O and
substituted with 0-3 R¹¹;

35 R¹¹ is independently selected at each occurrence from the
group: H, aryl substituted with 0-1 R¹², a 5-10 membered
heterocyclic ring system containing 1-4 heteroatoms

independently selected from N, S, and O and substituted with 0-1 R¹², C₃₋₁₀ cycloalkyl substituted with 0-1 R¹², polyalkylene glycol substituted with 0-1 R¹², carbohydrate substituted with 0-1 R¹², cyclodextrin substituted with 0-1 R¹², amino acid substituted with 0-1 R¹², polycarboxyalkyl substituted with 0-1 R¹², polyazaalkyl substituted with 0-1 R¹², peptide substituted with 0-1 R¹², wherein the peptide is comprised of 2-10 amino acids, and a bond to C_h;

R¹² is a bond to C_h;

k is selected from 0, 1, and 2;

h is selected from 0, 1, and 2;

h' is selected from 0, 1, 2, 3, 4, and 5;

h'' is selected from 0, 1, 2, 3, 4, and 5;

g is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g''' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

s is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

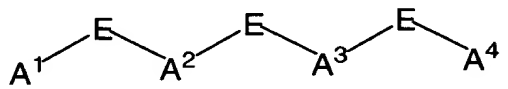
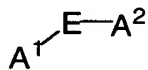
s' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

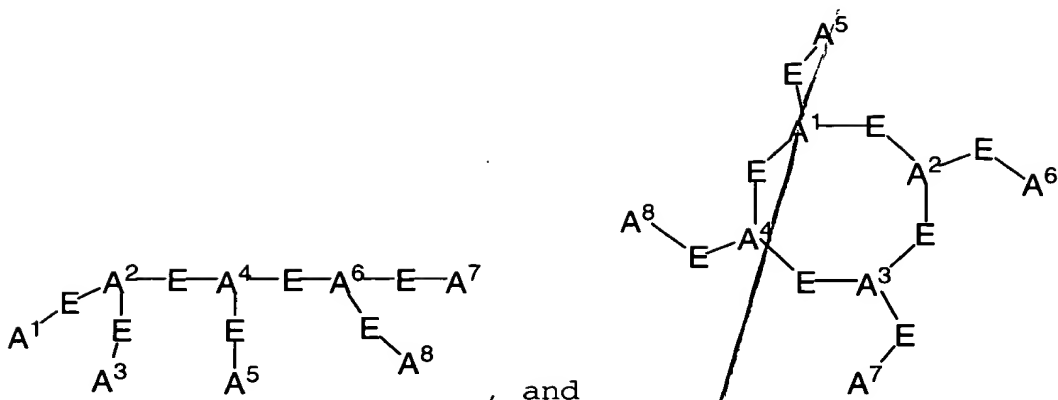
s'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

t is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

t' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

C_h is a metal bonding unit having a formula selected from the group:





A¹, A², A³, A⁴, A⁵, A⁶, A⁷, and A⁸ are independently selected at each occurrence from the group N, NR¹³, NR¹³R¹⁴, S, SH, S(Pg), O, OH, PR¹³, PR¹³R¹⁴, P(O)R¹⁵R¹⁶, and a bond to L_n;

E is a bond, CH, or a spacer group independently selected at each occurrence from the group: C₁-C₁₀ alkyl substituted with 0-3 R¹⁷, aryl substituted with 0-3 R¹⁷, C₃-₁₀ cycloalkyl substituted with 0-3 R¹⁷, heterocyclo-C₁-₁₀ alkyl substituted with 0-3 R¹⁷, wherein the heterocyclo group is a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O, C₆-₁₀ aryl-C₁-₁₀ alkyl substituted with 0-3 R¹⁷, C₁-₁₀ alkyl-C₆-₁₀ aryl- substituted with 0-3 R¹⁷, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R¹⁷;

R¹³, and R¹⁴ are each independently selected from the group: a bond to L_n, hydrogen, C₁-C₁₀ alkyl substituted with 0-3 R¹⁷, aryl substituted with 0-3 R¹⁷, C₁-₁₀ cycloalkyl substituted with 0-3 R¹⁷, heterocyclo-C₁-₁₀ alkyl substituted with 0-3 R¹⁷, wherein the heterocyclo group is a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O, C₆-₁₀ aryl-C₁-₁₀ alkyl substituted with 0-3 R¹⁷, C₁-₁₀ alkyl-C₆-₁₀ aryl- substituted with 0-3 R¹⁷, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and

546
37
cont

substituted with 0-3 R¹⁷, and an electron, provided that when one of R¹³ or R¹⁴ is an electron, then the other is also an electron;

5 alternatively, R¹³ and R¹⁴ combine to form =C(R²⁰)(R²¹);

15 R¹⁵ and R¹⁶ are each independently selected from the group: a
bond to L_n, -OH, C₁-C₁₀ alkyl substituted with 0-3 R¹⁷,
C₁-C₁₀ alkyl substituted with 0-3 R¹⁷, aryl substituted
10 with 0-3 R¹⁷, C₃-₁₀ cycloalkyl substituted with 0-3 R¹⁷,
heterocyclo-C₁-₁₀ alkyl substituted with 0-3 R¹⁷, wherein
the heterocyclo group is a 5-10 membered heterocyclic
ring system containing 1-4 heteroatoms independently
selected from N, S, and O, C₆-₁₀ aryl-C₁-₁₀ alkyl
substituted with 0-3 R¹⁷, C₁-₁₀ alkyl-C₆-₁₀ aryl-
substituted with 0-3 R¹⁷, and a 5-10 membered
heterocyclic ring system containing 1-4 heteroatoms
independently selected from N, S, and O and substituted
with 0-3 R¹⁷;

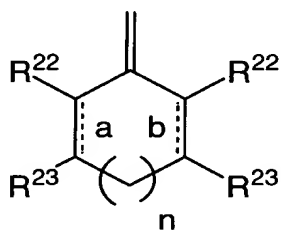
20 R¹⁷ is independently selected at each occurrence from the
group: a bond to L_n, =O, F, Cl, Br, I, -CF₃, -CN,
-CO₂R¹⁸, -C(=O)R¹⁸, -C(=O)N(R¹⁸)₂, -CHO, -CH₂OR¹⁸,
-OC(=O)R¹⁸, -OC(=O)OR^{18a}, -OR¹⁸, -OC(=O)N(R¹⁸)₂,
25 -NR¹⁹C(=O)R¹⁸, -NR¹⁹C(=O)OR^{18a}, -NR¹⁹C(=O)N(R¹⁸)₂,
-NR¹⁹SO₂N(R¹⁸)₂, -NR¹⁹SO₂R^{18a}, -SO₃H, -SO₂R^{18a}, -SR¹⁸,
-S(=O)R^{18a}, -SO₂N(R¹⁸)₂, -N(R¹⁸)₂, -NHC(=S)NHR¹⁸, =NOR¹⁸,
NO₂, -C(=O)NHR¹⁸, -C(=O)NHN(R¹⁸)₂, -OCH₂CO₂H,
2-(1-morpholino)ethoxy, C₁-C₅ alkyl, C₂-C₄ alkenyl, C₃-C₆
30 cycloalkyl, C₃-C₆ cycloalkylmethyl, C₂-C₆ alkoxyalkyl,
aryl substituted with 0-2 R¹⁸, and a 5-10 membered
heterocyclic ring system containing 1-4 heteroatoms
independently selected from N, S, and O;

35 R¹⁸, R^{18a}, and R¹⁹ are independently selected at each
occurrence from the group: a bond to L_n, H, C₁-C₆ alkyl,
phenyl, benzyl, C₁-C₆ alkoxy, halide, nitro, cyano, and
trifluoromethyl;

Pg is a thiol protecting group;

5 R^{20} and R^{21} are independently selected from the group: H,
C₁-C₁₀ alkyl, -CN, -CO₂R²⁵, -C(=O)R²⁵, -C(=O)N(R²⁵)₂,
C₂-C₁₀ 1-alkene substituted with 0-3 R²³, C₂-C₁₀ 1-alkyne
substituted with 0-3 R²³, aryl substituted with 0-3 R²³,
unsaturated 5-10 membered heterocyclic ring system
10 containing 1-4 heteroatoms independently selected from N,
S, and O and substituted with 0-3 R²³, and unsaturated
C₃-10 carbocycle substituted with 0-3 R²³;

alternatively, R^{20} and R^{21} , taken together with the divalent
carbon radical to which they are attached form:



20 R^{22} and R^{23} are independently selected from the group: H, R^{24} ,
C₁-C₁₀ alkyl substituted with 0-3 R^{24} , C₂-C₁₀ alkenyl
substituted with 0-3 R^{24} , C₂-C₁₀ alkynyl substituted with
0-3 R^{24} , aryl substituted with 0-3 R^{24} , a 5-10 membered
heterocyclic ring system containing 1-4 heteroatoms
independently selected from N, S, and O and substituted
with 0-3 R^{24} , and C₃-10 carbocycle substituted with 0-3
 R^{24} ;

25 alternatively, R^{22} , R^{23} taken together form a fused aromatic
or a 5-10 membered heterocyclic ring system containing
1-4 heteroatoms independently selected from N, S, and O;

30 **a** and **b** indicate the positions of optional double bonds and **n**
is 0 or 1;

5
10
15
20
25
30
35
660000 4243260

*Sub
39
done*

R^{24} is independently selected at each occurrence from the group: $=O$, F , Cl , Br , I , $-CF_3$, $-CN$, $-CO_2R^{25}$, $-C(=O)R^{25}$, $-C(=O)N(R^{25})_2$, $-N(R^{25})_3^+$, $-CH_2OR^{25}$, $-OC(=O)R^{25}$, $-OC(=O)OR^{25a}$, $-OR^{25}$, $-OC(=O)N(R^{25})_2$, $-NR^{26}C(=O)R^{25}$, $-NR^{26}C(=O)OR^{25a}$, $-NR^{26}C(=O)N(R^{25})_2$, $-NR^{26}SO_2N(R^{25})_2$, $-NR^{26}SO_2R^{25a}$, $-SO_3H$, $-SO_2R^{25a}$, $-SR^{25}$, $-S(=O)R^{25a}$, $-SO_2N(R^{25})_2$, $-N(R^{25})_2$, $=NOR^{25}$, $-C(=O)NHOR^{25}$, $-OCH_2CO_2H$, and 2-(1-morpholino)ethoxy; and,

10 R^{25} , R^{25a} , and R^{26} are each independently selected at each occurrence from the group: hydrogen and C_1 - C_6 alkyl;

and a pharmaceutically acceptable salt thereof.

15 4. A compound according to Claim 3, the present invention provides a compound, wherein:

20 L is glycine;

25 R^1 is an amino acid, optionally substituted with a bond to L_n , independently selected at each occurrence from the group: L-valine, D-valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, tyrosine, phenylalanine, phenylglycine, cyclohexylalanine, homophenylalanine, lysine, ornithine, 1,2-diaminobutyric acid, and 1,2-diaminopropionic acid;

30 R^2 is an amino acid, optionally substituted with a bond to L_n , independently selected at each occurrence from the group: valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, tyrosine, L-phenylalanine, D-phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine, homophenylalanine, 35 L-1-naphthylalanine, D-1-naphthylalanine, lysine, ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, and 2-aminothiazole-4-acetic acid;

membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1 R¹⁰;

5 R⁶, R^{6a}, R⁷, R^{7a}, R⁸, R^{8a}, R⁹, and R^{9a} are independently selected at each occurrence from the group: H, =O, COOH, SO₃H, C₁-C₅ alkyl substituted with 0-1 R¹⁰, aryl substituted with 0-1 R¹⁰, benzyl substituted with 0-1 R¹⁰, and C₁-C₅ alkoxy substituted with 0-1 R¹⁰,
10 NHC(=O)R¹¹, C(=O)NHR¹¹, NHC(=O)NHR¹¹, NHR¹¹, R¹¹, and a bond to C_h;

R¹⁰ is independently selected at each occurrence from the group: COOR¹¹, OH, NHR¹¹, SO₃H, aryl substituted with
15 0-1 R¹¹, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-1 R¹¹, C₁-C₅ alkyl substituted with 0-1 R¹², C₁-C₅ alkoxy substituted with 0-1 R¹², and a bond to C_h;

20 R¹¹ is independently selected at each occurrence from the group: H, aryl substituted with 0-1 R¹², a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted
25 with 0-1 R¹², polyalkylene glycol substituted with 0-1 R¹², carbohydrate substituted with 0-1 R¹², cyclodextrin substituted with 0-1 R¹², amino acid substituted with 0-1 R¹², and a bond to C_h;

30 k is 0 or 1;
h is 0 or 1;
h' is 0 or 1;
s is selected from 0, 1, 2, 3, 4, and 5;
s' is selected from 0, 1, 2, 3, 4, and 5;
35 s" is selected from 0, 1, 2, 3, 4, and 5;
t is selected from 0, 1, 2, 3, 4, and 5;

A¹, A², A³, A⁴, A⁵, A⁶, A⁷, and A⁸ are independently selected at each occurrence from the group: NR¹³, NR¹³R¹⁴, S, SH, S(Pg), OH, and a bond to L_n;

5 E is a bond, CH, or a spacer group independently selected at each occurrence from the group: C₁-C₁₀ alkyl substituted with 0-3 R¹⁷, aryl substituted with 0-3 R¹⁷, C₃₋₁₀ cycloalkyl substituted with 0-3 R¹⁷, and a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms
10 independently selected from N, S, and O and substituted with 0-3 R¹⁷;

R¹³, and R¹⁴ are each independently selected from the group:
15 a bond to L_n, hydrogen, C₁-C₁₀ alkyl substituted with 0-3 R¹⁷, aryl substituted with 0-3 R¹⁷, a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R¹⁷, and an electron, provided that when one of R¹³ or R¹⁴ is an electron, then the other is also an
20 electron;

alternatively, R¹³ and R¹⁴ combine to form =C(R²⁰)(R²¹);

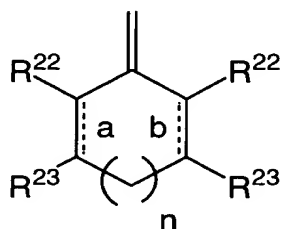
R¹⁷ is independently selected at each occurrence from the
25 group: a bond to L_n, =O, F, Cl, Br, I, -CF₃, -CN, -CO₂R¹⁸, -C(=O)R¹⁸, -C(=O)N(R¹⁸)₂, -CH₂OR¹⁸, -OC(=O)R¹⁸, -OC(=O)OR^{18a}, -OR¹⁸, -OC(=O)N(R¹⁸)₂, -NR¹⁹C(=O)R¹⁸, -NR¹⁹C(=O)OR^{18a}, -NR¹⁹C(=O)N(R¹⁸)₂, -NR¹⁹SO₂N(R¹⁸)₂, -NR¹⁹SO₂R^{18a}, -SO₃H, -SO₂R^{18a}, -S(=O)R^{18a}, -SO₂N(R¹⁸)₂,
30 -N(R¹⁸)₂, -NHC(=S)NHR¹⁸, =NOR¹⁸, -C(=O)NHN(R¹⁸)₂, -OCH₂CO₂H, and 2-(1-morpholino)ethoxy;

R¹⁸, R^{18a}, and R¹⁹ are independently selected at each
35 occurrence from the group: a bond to L_n, H, and C₁-C₆ alkyl;

R²⁰ and R²¹ are independently selected from the group: H, C₁-C₅ alkyl, -CO₂R²⁵, C₂-C₅ 1-alkene substituted with 0-3

R²³, C₂-C₅ 1-alkyne substituted with 0-3 R²³, aryl substituted with 0-3 R²³, and unsaturated 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O and substituted with 0-3 R²³;

alternatively, R²⁰ and R²¹, taken together with the divalent carbon radical to which they are attached form:



R²² and R²³ are independently selected from the group: H, and R²⁴;

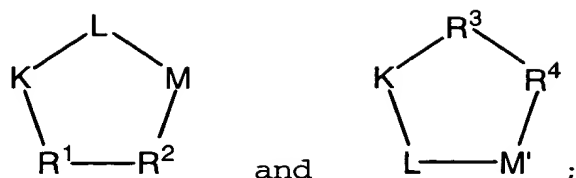
alternatively, R²², R²³ taken together form a fused aromatic or a 5-10 membered heterocyclic ring system containing 1-4 heteroatoms independently selected from N, S, and O;

R²⁴ is independently selected at each occurrence from the group: -CO₂R²⁵, -C(=O)N(R²⁵)₂, -CH₂OR²⁵, -OC(=O)R²⁵, -OR²⁵, -SO₃H, -N(R²⁵)₂, and -OCH₂CO₂H; and,

R²⁵ is independently selected at each occurrence from the group: H and C₁-C₃ alkyl.

5. A compound according to Claim 4, the present invention provides a compound, wherein:

Q is a peptide selected from the group:



R^1 is L-valine, D-valine, D-lysine optionally substituted on
 the ϵ amino group with a bond to L_n or L-lysine
 5 optionally substituted on the ϵ amino group with a bond
 to L_n ;

R^2 is L-phenylalanine, D-phenylalanine, D-1-naphthylalanine,
 2-aminothiazole-4-acetic acid, L-lysine optionally
 10 substituted on the ϵ amino group with a bond to L_n or
 tyrosine, the tyrosine optionally substituted on the
 hydroxy group with a bond to L_n ;

R^3 is D-valine, D-phenylalanine, or L-lysine optionally
 15 substituted on the ϵ amino group with a bond to L_n ;

R^4 is D-phenylalanine, D-tyrosine substituted on the hydroxy
 group with a bond to L_n , or L-lysine optionally
 substituted on the ϵ amino group with a bond to L_n ;

20 provided that one of R^1 and R^2 in each Q is substituted with a
 bond to L_n , and further provided that when R^2 is
 2-aminothiazole-4-acetic acid, K is N-methylarginine;

25 d is 1 or 2;

W is independently selected at each occurrence from the group:
 $NHC(=O)$, $C(=O)NH$, $C(=O)$, $(CH_2CH_2O)_s$, and $(CH_2CH_2CH_2O)_t$;

30 R^6 , R^{6a} , R^7 , R^{7a} , R^8 , R^{8a} , R^9 , and R^{9a} are independently
 selected at each occurrence from the group: H ,
 $NHC(=O)R^{11}$, and a bond to C_h ;

k is 0;

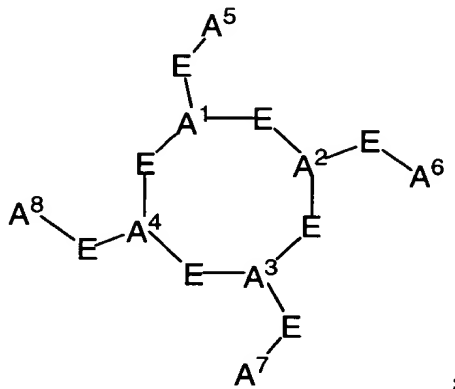


R¹⁸ is a bond to L_n;

R²⁴ is selected from the group: -CO₂R²⁵, -OR²⁵, -SO₃H, and -N(R²⁵)₂;

5

R²⁵ is independently selected at each occurrence from the group: hydrogen and methyl;



alternatively, C_h is

10

A¹, A², A³, and A⁴ are each N;

A⁵, A⁶, and A⁸ are each OH;

15

A⁷ is a bond to L_n;

E is a C₂ alkyl substituted with 0-1 R¹⁷; and,

R¹⁷ is =O.

20

6. A compound according to Claim 3, the present invention provides a compound selected from the group:

25

(a) cyclo{Arg-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};

30

(b) cyclo{Arg-Gly-Asp-D-Tyr((N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-18-

- (w) cyclo{Lys-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};
- 5 (x) cyclo{Cys(2-aminoethyl)-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};
- 10 (y) cyclo{HomoLys-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};
- 15 (z) cyclo{Orn(d-N-Benzylcarbamoyl)-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};
- 20 (aa) cyclo{Dap(b-(2-benzimidazolylacetyl))-Gly-Asp-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-Val};
- 25 (bb) cyclo{Orn(d-N-2-ImidazolinyI)-Gly-Asp-D-Phe-Lys(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]]);
- 30 (cc) cyclo{Orn(d-N-Benzylcarbamoyl)-Gly-Asp-D-Phe-Lys(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]]);
- 35 (dd) cyclo{Lys-D-Val-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-D-Asp-Gly};
- (ee) cyclo{Orn(d-N-Benzylcarbamoyl)-D-Val-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridinyl]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-D-Asp-Gly}; and,

(ff) cyclo{Orn(d-N-2-Imidazoliny1)-D-Val-D-Tyr(N-[2-[[[5-[carbonyl]-2-pyridiny1]hydrazono]methyl]-benzenesulfonic acid]-3-aminopropyl)-D-Asp-Gly};

5 or a pharmaceutically acceptable salt form thereof.

7. A kit comprising a compound of Claim 3, or a pharmaceutically acceptable salt form thereof and a
10 pharmaceutically acceptable carrier.

8. A kit according to Claim 7, wherein the kit further comprises one or more ancillary ligands and a reducing agent.
15

9. A kit according to Claim 8, wherein the ancillary ligands are tricine and TPPTS.
20

10. A kit according to Claim 9, wherein the reducing agent is tin(II).
25

11. A diagnostic or therapeutic metallopharmaceutical composition, comprising: a metal, a chelator capable of chelating the metal and a targeting moiety, wherein the targeting moiety is bound to the chelator, is a peptide or peptidomimetic and binds to a receptor that is upregulated during angiogenesis and the compound has 0-1 linking groups between the targeting moiety and chelator.
30

12. A composition according to Claim 11, wherein the
35 metallopharmaceutical is a diagnostic radiopharmaceutical, the metal is a radioisotope selected from the group: ^{99m}Tc , ^{95}Tc , ^{111}In , ^{62}Cu , ^{64}Cu , ^{67}Ga , and ^{68}Ga , the targeting moiety is a peptide or a mimetic thereof and the receptor is selected from

the group: EGFR, FGFR, PDGFR, Flk-1/KDR, Flt-1, Tek, Tie, neuropilin-1, endoglin, endosialin, Axl, $\alpha_v\beta_3$, $\alpha_v\beta_5$, $\alpha_5\beta_1$, $\alpha_4\beta_1$, $\alpha_1\beta_1$, and $\alpha_2\beta_2$ and the linking group is present between the targeting moiety and chelator.

13. A composition according to Claim 12, wherein the targeting moiety is a cyclic pentapeptide and the receptor is $\alpha_v\beta_3$.

14. A composition according to Claim 13, wherein the radioisotope is ^{99m}Tc or ^{95}Tc , the radiopharmaceutical further comprises a first ancillary ligand and a second ancillary ligand capable of stabilizing the radiopharmaceutical.

15. A composition according to Claim 14, wherein the radioisotope is ^{99m}Tc .

16. A composition according to Claim 15, wherein the radiopharmaceutical is selected from the group:

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-Gly-Asp-D-Tyr}(\text{N}-[[5-[\text{carbonyl}]-2\text{-pyridinyl}]\text{diazenido}]-3\text{-aminopropyl})-\text{Val}));$

$^{99m}\text{Tc}(\text{tricine})(\text{TPPMS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5-[\text{carbonyl}]-2\text{-pyridinyl}]\text{diazenido}]-3\text{-aminopropyl})-\text{D-Asp-Gly}));$

$^{99m}\text{Tc}(\text{tricine})(\text{TPPDS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5-[\text{carbonyl}]-2\text{-pyridinyl}]\text{diazenido}]-3\text{-aminopropyl})-\text{D-Asp-Gly}));$

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-D-Val-D-Tyr}(\text{N}-[[5-[\text{carbonyl}]-2\text{-pyridinyl}]\text{diazenido}]-3\text{-aminopropyl})-\text{D-Asp-Gly}));$

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{Arg-Gly-Asp-D-Phe-Lys}(\text{N}-[[5-[\text{carbonyl}]-2\text{-pyridinyl}]\text{diazenido}]));$

Glu(cyclo{D-Lys-D-Phe-D-Asp-Gly-Arg})-cyclo{D-Lys-D-Phe-D-Asp-Gly-Arg});

5 $^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}\{\text{D-Phe-D-Lys}([2-[[[5-[\text{carbonyl}]-2-\text{pyridinyl}]\text{hydrazono}]\text{methyl}]-\text{benzenesulfonic acid}])-\text{D-Asp-Gly-Arg})\});$

10 $^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}(\text{N-Me-Arg-Gly-Asp-ATA-D-Lys}(\text{N}-[[5-[\text{carbonyl}]-2-\text{pyridinyl}]\text{diazenido}])))$;

$^{99m}\text{Tc}(\text{tricine})(\text{TPPTS})(\text{cyclo}\{\text{Cit-Gly-Asp-D-Phe-Lys}([2-[[[5-[\text{carbonyl}]-2-\text{pyridinyl}]\text{hydrazono}]\text{methyl}]-\text{benzenesulfonic acid}])))$; and,

15 $^{99m}\text{Tc}(\text{tricine})(1,2,4\text{-triazole})(\text{cyclo}(\text{Arg-Gly-Asp-D-Tyr}(\text{N}-[[5-[\text{carbonyl}]-2-\text{pyridinyl}]\text{diazenido}]-3\text{-aminopropyl})-\text{Val}))$.

20 17. A composition according to Claim 13, wherein the radioisotope is ^{111}In .

25 18. A composition according to Claim 17, wherein the radiopharmaceutical is selected from the group:

(DOTA- ^{111}In)-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe};

cyclo(Arg-Gly-Asp-D-Phe-Lys(DTPA- ^{111}In)); and,

30 cyclo(Arg-Gly-Asp-D-Phe-Lys) $_2$ (DTPA- ^{111}In).

35 19. A composition according to Claim 11, wherein the metallopharmaceutical is a therapeutic radiopharmaceutical, the metal is a radioisotope selected from the group: ^{186}Re , ^{188}Re , ^{153}Sm , ^{166}Ho , ^{177}Lu , ^{149}Pm , ^{90}Y , ^{212}Bi , ^{103}Pd , ^{109}Pd , ^{159}Gd , ^{140}La , ^{198}Au , ^{199}Au , ^{169}Yb , ^{175}Yb , ^{165}Dy , ^{166}Dy , ^{67}Cu ,

105Rh, 111Ag, and 192Ir, the targeting moiety is a peptide or a mimetic thereof and the receptor is selected from the group: EGFR, FGFR, PDGFR, Flk-1/KDR, Flt-1, Tek, Tie, neuropilin-1, endoglin, endosialin, Axl, $\alpha_v\beta_3$, $\alpha_v\beta_5$, $\alpha_5\beta_1$, $\alpha_4\beta_1$, $\alpha_1\beta_1$, and $\alpha_2\beta_2$ and the linking group is present between the targeting moiety and chelator.

20. A composition according to Claim 19, wherein the targeting moiety is a cyclic pentapeptide and the receptor is $\alpha_v\beta_3$.

21. A composition according to Claim 20, wherein the radioisotope is ^{153}Sm .

22. A composition according to Claim 21, wherein the radiopharmaceutical is selected from the group:

cyclo(Arg-Gly-Asp-D-Phe-Lys(DTPA- ^{153}Sm));

cyclo(Arg-Gly-Asp-D-Phe-Lys)₂(DTPA- ^{153}Sm); and,

cyclo(Arg-Gly-Asp-D-Tyr(N-DTPA(^{153}Sm)-3-aminopropyl)-Val).

23. A composition according to Claim 20, wherein the radioisotope is ^{177}Lu .

24. A composition according to Claim 23, wherein the radiopharmaceutical is selected from the group:

cyclo(Arg-Gly-Asp-D-Phe-Lys(DTPA- ^{177}Lu));

(DOTA- ^{177}Lu)-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe};

cyclo(Arg-Gly-Asp-D-Phe-Lys)₂(DTPA-¹⁷⁷Lu); and,

cyclo(Arg-Gly-Asp-D-Tyr(N-DTPA(¹⁷⁷Lu)-3-aminopropyl)-Val).

5

25. A composition according to Claim 20, wherein the radioisotope is ⁹⁰Y.

10

26. A composition according to Claim 25, wherein the radiopharmaceutical is:

(DOTA-⁹⁰Y)-Glu(cyclo{Lys-Arg-Gly-Asp-D-Phe})-cyclo{Lys-Arg-Gly-Asp-D-Phe};

15

27. A composition according to Claim 11, wherein the metallopharmaceutical is a MRI contrast agent, the metal is a paramagnetic metal ion selected from the group: Gd(III), Dy(III), Fe(III), and Mn(II), the targeting moiety is a peptide or a mimetic thereof and the receptor is selected from the group: EGFR, FGFR, PDGFR, Flk-1/KDR, Flt-1, Tek, Tie, neuropilin-1, endoglin, endosialin, Axl, $\alpha_v\beta_3$, $\alpha_v\beta_5$, $\alpha_5\beta_1$, $\alpha_4\beta_1$, $\alpha_1\beta_1$, and $\alpha_2\beta_2$ and the linking group is present between the targeting moiety and chelator.

20

25

28. A composition according to Claim 27, wherein the targeting moiety is a cyclic pentapeptide and the receptor is $\alpha_v\beta_3$.

30

29. A composition according to Claim 28, wherein the metal ion is Gd(III).

35

30. A composition according to Claim 29, wherein the contrast agent is:

cyclo(Arg-Gly-Asp-D-Tyr(N-DTPA(Gd(III))-3-aminopropyl)-Val).

5

516
35
10
31. A composition according to Claim 11, wherein the metallopharmaceutical is a X-ray contrast agent, the metal is selected from the group: Re, Sm, Ho, Lu, Pm, Y, Bi, Pd, Gd, La, Au, Au, Yb, Dy, Cu, Rh, Ag, and Ir, the targeting moiety is a cyclic pentapeptide, the receptor is $\alpha_v\beta_3$, and the linking group is present between the targeting moiety and chelator.

15

32. A method of treating rheumatoid arthritis in a patient comprising: administering a therapeutic radiopharmaceutical of Claim 11 capable of localizing in new angiogenic vasculature to a patient by injection or infusion.

20

33. A method of treating cancer in a patient comprising: administering to a patient in need thereof a therapeutic radiopharmaceutical of Claim 11 by injection or infusion.

25

34. A method of imaging formation of new blood vessels in a patient comprising: (1) administering a diagnostic radiopharmaceutical, a MRI contrast agent, or a X-ray contrast agent of of Claim 11 to a patient by injection or infusion; (2) imaging the area of the patient wherein the desired formation of new blood vessels is located.

30

35. A method of imaging cancer in a patient comprising: (1) administering a diagnostic radiopharmaceutical of Claim 12 to a patient by injection or infusion; (2) imaging the patient

35

acid, L-tyrosine, L-phenylalanine, L-thienylalanine,
L-phenylglycine, L-cyclohexylalanine,
L-homophenylalanine, L-1-naphthylalanine, L-lysine,
L-serine, L-ornithine, L-1,2-diaminobutyric acid,
5 L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine,
L-methionine, and 2-aminothiazole-4-acetic acid;

provided that one of R^1 , R^2 , R^3 , R^4 , and R^5 in each Q is
substituted with a bond to L_n , further provided that when
10 R^2 is 2-aminothiazole-4-acetic acid, K is
N-methylarginine, further provided that when R^4 is
2-aminothiazole-4-acetic acid, K and K' are
N-methylarginine, and still further provided that when R^5
is 2-aminothiazole-4-acetic acid, K' is N-methylarginine;

15 d is selected from 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

S_f is a surfactant which is a lipid or a compound of the

formula: $A^9 - E^1 - A^{10}$;

20 A^9 is selected from the group: OH and OR^{27} ;

A^{10} is OR^{27} ;

25 R^{27} is $C(=O)C_{1-20}$ alkyl;

E^1 is C_{1-10} alkylene substituted with 1-3 R^{28} ;

30 R^{28} is independently selected at each occurrence from the
group: R^{30} , $-PO_3H-R^{30}$, $=O$, $-CO_2R^{29}$, $-C(=O)R^{29}$,
 $-C(=O)N(R^{29})_2$, $-CH_2OR^{29}$, $-OR^{29}$, $-N(R^{29})_2$, C_1-C_5 alkyl,
and C_2-C_4 alkenyl;

35 R^{29} is independently selected at each occurrence from the
group: R^{30} , H, C_1-C_6 alkyl, phenyl, benzyl, and
trifluoromethyl;

660EEQ" 42478260

R³⁰ is a bond to L_n;

L_n is a linking group having the formula:

5 (CR⁶R⁷)_g-(W)_h-(CR^{6a}R^{7a})_{g'}-(Z)_k-(W)_{h'}-(CR⁸R⁹)_{g''}-(W)_{h''}-(CR^{8a}R^{9a})_{g'''}.

W is independently selected at each occurrence from the group:

O, S, NH, NHC(=O), C(=O)NH, C(=O), C(=O)O, OC(=O),
NHC(=S)NH, NHC(=O)NH, SO₂, (OCH₂CH₂)₂₀₋₂₀₀, (CH₂CH₂O)₂₀₋
10 200, (OCH₂CH₂CH₂)₂₀₋₂₀₀, (CH₂CH₂CH₂O)₂₀₋₂₀₀, and (aa)_t;

aa is independently at each occurrence an amino acid;

15 Z is selected from the group: aryl substituted with 0-3 R¹⁰,
C₃₋₁₀ cycloalkyl substituted with 0-3 R¹⁰, and a 5-10
membered heterocyclic ring system containing 1-4
heteroatoms independently selected from N, S, and O and
substituted with 0-3 R¹⁰;

20 R⁶, R^{6a}, R⁷, R^{7a}, R⁸, R^{8a}, R⁹ and R^{9a} are independently selected
at each occurrence from the group: H, =O, COOH, SO₃H,
PO₃H, C₁₋₅ alkyl substituted with 0-3 R¹⁰, aryl
substituted with 0-3 R¹⁰, benzyl substituted with 0-3
R¹⁰, and C₁₋₅ alkoxy substituted with 0-3 R¹⁰,
25 NHC(=O)R¹¹, C(=O)NHR¹¹, NHC(=O)NHR¹¹, NHR¹¹, R¹¹, and a
bond to S_f;

R¹⁰ is independently selected at each occurrence from the
group: a bond to S_f, COOR¹¹, OH, NHR¹¹, SO₃H, PO₃H, aryl
30 substituted with 0-3 R¹¹, C₁₋₅ alkyl substituted with 0-1
R¹², C₁₋₅ alkoxy substituted with 0-1 R¹², and a 5-10
membered heterocyclic ring system containing 1-4
heteroatoms independently selected from N, S, and O and
substituted with 0-3 R¹¹;

35 R¹¹ is independently selected at each occurrence from the
group: H, aryl substituted with 0-1 R¹², a 5-10 membered
heterocyclic ring system containing 1-4 heteroatoms

independently selected from N, S, and O and substituted with 0-1 R¹², C₃₋₁₀ cycloalkyl substituted with 0-1 R¹², amino acid substituted with 0-1 R¹², and a bond to S_f;

5 R¹² is a bond to S_f;

k is selected from 0, 1, and 2;

h is selected from 0, 1, and 2;

h' is selected from 0, 1, 2, 3, 4, and 5;

10 h'' is selected from 0, 1, 2, 3, 4, and 5;

g is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g'' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

g''' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

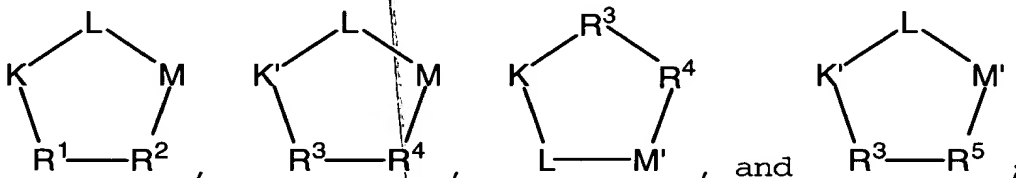
15 t' is selected from 0, 1, 2, 3, 4, 5, 6, 7, 8, 9, and 10;

and a pharmaceutically acceptable salt thereof.

20 41. A compound according to Claim 40, wherein the compound is of the formula:



wherein, Q is a cyclic pentapeptide independently selected from the group:



30 K is an L-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine, δ -N-2-imidazolinylnornithine, δ -N-benzylcarbamoylnornithine, and β -2-benzimidazolylacetyl-1,2-diaminopropionic acid;

35

K' is a D-amino acid independently selected at each occurrence from the group: arginine, citrulline, N-methylarginine, lysine, homolysine, 2-aminoethylcysteine, δ -N-2-imidazolinylnornithine, δ -N-benzylcarbamoylnornithine, and β -2-benzimidazolylacetyl-1,2-diaminopropionic acid;

L is independently selected at each occurrence from the group: glycine, L-alanine, and D-alanine;

M is L-aspartic acid;

M' is D-aspartic acid;

R¹ is an amino acid substituted with 0-1 bonds to L_n, independently selected at each occurrence from the group: glycine, L-valine, D-valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, phenylalanine, thienylalanine, phenylglycine, cyclohexylalanine, homophenylalanine, 1-naphthylalanine, lysine, serine, ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, cysteine, penicillamine, and methionine;

R² is an amino acid, substituted with 0-1 bonds to L_n, independently selected at each occurrence from the group: glycine, valine, alanine, leucine, isoleucine, norleucine, 2-aminobutyric acid, 2-aminohexanoic acid, tyrosine, L-phenylalanine, D-phenylalanine, thienylalanine, phenylglycine, biphenylglycine, cyclohexylalanine, homophenylalanine, L-1-naphthylalanine, D-1-naphthylalanine, lysine, serine, ornithine, 1,2-diaminobutyric acid, 1,2-diaminopropionic acid, cysteine, penicillamine, methionine, and 2-aminothiazole-4-acetic acid;

R³ is an amino acid, substituted with 0-1 bonds to L_n, independently selected at each occurrence from the group:

glycine, D-valine, D-alanine, D-leucine, D-isoleucine, D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic acid, D-tyrosine, D-phenylalanine, D-thienylalanine, D-phenylglycine, D-cyclohexylalanine, D-homophenylalanine, D-1-naphthylalanine, D-lysine, D-serine, D-ornithine, D-1,2-diaminobutyric acid, D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine, and D-methionine;

10 R⁴ is an amino acid, substituted with 0-1 bonds to L_n,
independently selected at each occurrence from the group:
glycine, D-valine, D-alanine, D-leucine, D-isoleucine,
D-norleucine, D-2-aminobutyric acid, D-2-aminohexanoic
acid, D-tyrosine, D-phenylalanine, D-thienylalanine,
15 D-phenylglycine, D-cyclohexylalanine,
D-homophenylalanine, D-1-naphthylalanine, D-lysine,
D-serine, D-ornithine, D-1,2-diaminobutyric acid,
D-1,2-diaminopropionic acid, D-cysteine, D-penicillamine,
D-methionine, and 2-aminothiazole-4-acetic acid;

R⁵ is an amino acid, substituted with 0-1 bonds to L_n, independently selected at each occurrence from the group: glycine, L-valine, L-alanine, L-leucine, L-isoleucine, L-norleucine, L-2-aminobutyric acid, L-2-aminohexanoic acid, L-tyrosine, L-phenylalanine, L-thienylalanine, L-phenylglycine, L-cyclohexylalanine, L-homophenylalanine, L-1-naphthylalanine, L-lysine, L-serine, L-ornithine, L-1,2-diaminobutyric acid, L-1,2-diaminopropionic acid, L-cysteine, L-penicillamine, L-methionine, and 2-aminothiazole-4-acetic acid;

provided that one of R¹, R², R³, R⁴, and R⁵ in each Q is substituted with a bond to L_n, further provided that when R² is 2-aminothiazole-4-acetic acid, K is N-methylarginine, further provided that when R⁴ is 2-aminothiazole-4-acetic acid, K and K' are N-methylarginine, and still further provided that when R⁵ is 2-aminothiazole-4-acetic acid, K' is N-methylarginine.

660000" 4478260

5 R⁶, R^{6a}, R⁷, R^{7a}, R⁸, R^{8a}, R⁹ and R^{9a} are independently selected at each occurrence from the group: H, =O, C₁-C₅ alkyl substituted with 0-3 R¹⁰, and C₁-C₅ alkoxy substituted with 0-3 R¹⁰, and a bond to S_f;

10 R¹⁰ is independently selected at each occurrence from the group: a bond to S_f, COOR¹¹, OH, NHR¹¹, C₁-5 alkyl substituted with 0-1 R¹², and C₁-5 alkoxy substituted with 0-1 R¹²;

15 R¹¹ is independently selected at each occurrence from the group: H, aryl substituted with 0-1 R¹², C₃-10 cycloalkyl substituted with 0-1 R¹², amino acid substituted with 0-1 R¹², and a bond to S_f;

R¹² is a bond to S_f;

20 k is selected from 0, 1, and 2;
h is selected from 0, 1, and 2;
h' is selected from 0, 1, 2, 3, 4, and 5;
h" is selected from 0, 1, 2, 3, 4, and 5;
g is selected from 0, 1, 2, 3, 4, and 5;
g' is selected from 0, 1, 2, 3, 4, and 5;
25 g" is selected from 0, 1, 2, 3, 4, and 5;
g'" is selected from 0, 1, 2, 3, 4, and 5;
s is selected from 0, 1, 2, 3, 4, and 5;
s' is selected from 0, 1, 2, 3, 4, and 5;
s" is selected from 0, 1, 2, 3, 4, and 5;
30 t is selected from 0, 1, 2, 3, 4, and 5;
t' is selected from 0, 1, 2, 3, 4, and 5;

and a pharmaceutically acceptable salt thereof.

35

42. A compound according to Claim 41, wherein the present invention provides a compound selected from the group:

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-(cyclo(Arg-Gly-Asp-D-Phe-Lys)-dodecane-1,12-dione;

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-((ω -amino-PEG₃₄₀₀- α -carbonyl)-cyclo(Arg-Gly-Asp-D-Phe-Lys))-dodecane-1,12-dione; and,

1-(1,2-Dipalmitoyl-sn-glycero-3-phosphoethanolamino)-12-((ω -amino-PEG₃₄₀₀- α -carbonyl)-Glu-(cyclo(Arg-Gly-Asp-D-Phe-Lys))₂)-Dodecane-1,12-dione.

43. An ultrasound contrast agent composition, comprising:

(a) a compound of Claim 40, comprising: a cyclic pentapeptide that binds to the integrin $\alpha_v\beta_3$, a surfactant and a linking group between the cyclic pentapeptide and the surfactant;

(b) a parenterally acceptable carrier; and,
(c) an echogenic gas.

44. An ultrasound contrast agent composition, further comprising: 1,2-dipalmitoyl-sn-glycero-3-phosphotidic acid, 1,2-dipalmitoyl-sn-glycero-3-phosphatidylcholine, and N-(methoxypolyethylene glycol 5000 carbamoyl)-1,2-dipalmitoyl-sn-glycero-3-phosphatidylethanolamine.

45. An ultrasound contrast agent composition, wherein, the echogenic gas is a C₂₋₅ perfluorocarbon.

46. A method of imaging cancer in a patient comprising: (1) administering, by injection or infusion, a ultrasound contrast agent composition of Claim 40 to a patient; and (2) imaging the patient using sonography.

47. A method of imaging formation of new blood vessels in a patient comprising: (1) administering, by injection or infusion, a ultrasound contrast agent composition of of Claim 5 40 to a patient; (2) imaging the area of the patient wherein the desired formation of new blood vessels is located.

48. A therapeutic radiopharmaceutical composition, 10 comprising:
(a) a therapeutic radiopharmaceutical of Claim 11; and,
(b) a parenterally acceptable carrier.

49. A diagnostic radiopharmaceutical composition, 15 comprising:
(a) a diagnostic radiopharmaceutical, a MRI contrast agent, or a X-ray contrast agent of Claim 11; and,
(b) a parenterally acceptable carrier. 20

50. A therapeutic radiopharmaceutical composition, comprising: a radiolabelled targeting moiety, wherein the targeting moiety is a compound Q of Claim 3 and the radiolabel 25 is a therapeutic isotope selected from the group: ^{35}S , ^{32}P , ^{125}I , ^{131}I , and ^{211}At .

51. A therapeutic radiopharmaceutical composition, 30 comprising: a radiolabelled targeting moiety, wherein the targeting moiety is a compound Q of Claim 5 and the radiolabel is a therapeutic isotope which is ^{131}I .